

#8

PCT09

## RAW SEQUENCE LISTING

DATE: 08/14/2001

PATENT APPLICATION: US/09/890,782

TIME: 10:55:50

Input Set : A:\PTO\_VSK.txt

Output Set: N:\CRF3\08142001\I890782.raw

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3 <110> APPLICANT: Rijksuniversiteit Leiden  
 5 <120> TITLE OF INVENTION: method of modulating metabolite biosynthesis in  
 6 recombinant cells  
 8 <130> FILE REFERENCE: BO 43339  
 ofc--> 10 <140> CURRENT APPLICATION NUMBER: US/09/890,782  
 c--> 11 <141> CURRENT FILING DATE: 2001-08-06  
 13 <160> NUMBER OF SEQ ID NOS: 21  
 15 <170> SOFTWARE: PatentIn Ver. 2.1  
 17 <210> SEQ ID NO: 1  
 18 <211> LENGTH: 1754  
 19 <212> TYPE: DNA  
 20 <213> ORGANISM: Catharanthus roseus  
 22 <400> SEQUENCE: 1  
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 24 ggaattacta aaatcggaag aagaaatcaa cgcgacgaaa gagaaaaaga acaaaagggt 120  
 25 ttcgtttttg taaagtttga ttcttgccgg agattttcga caaaggagtg ggcaatttgt 180  
 26 gcaatacttc tgagaaaaatt gaaagagata caaggatggc tcttcttgat caggcatcca 240  
 27 atttgagtc catgcctttt gatttcaacta gaaagaggaa gtcgaggagg agggatggt 300  
 28 ctaagaacgt agcggagaca cttgcaaagt ggaaagagta taatgagaaa cttgatgctt 360  
 29 tagatggagg gaaaccagct cggaagggtc ctgccaaaag atcaaaaaag ggatgtatga 420  
 30 aaggtaaaag aggccctgag aattctcact gcaaatacag aggagttagg cagaggacat 480  
 31 ggggtaaatg ggtggccgaa attcgggaac caaacagggg tagcaggctt tggttgggta 540  
 32 cattcagaaa cgcgatatga gctgcacttg cttatgatga agcagcgagg gcgatgtatg 600  
 33 ggccttgtgc taggcttaat cttccgaact atagggcttc agaagaatct tcttccttgc 660  
 34 caacaacatc aggatcagat acgactactg cttctggcat ctccagaggtc tctgtctatg 720  
 35 aagacaaaaa gttcacacca gttgtttccg gattgaaaca agatgacaag ggtgaatcat 780  
 36 tagagtcagc tgatagtaaa cctcaactcc tggctcgatgc tggcactccc atgagtgcag 840  
 37 tgaagggaaga accaaaagaa tatcaggtta tggattccca gtctgaaggg caattcggag 900  
 38 acgaggaacc gcctagcaag cttgtttgta aagaagtcga ctttgggcag gatcaagctg 960  
 39 ttgttcctgc tgtaaaaaat gctgaggaga tgggtggaga gatgggtgga gatatactga 1020  
 40 aaggctgttc tttgtctgag atgtttgatg tggacgagtt gttgagcgtt ttagattcta 1080  
 41 caccctcca tgcctcagat ttccagcatg gcatgggaaa tggtaatgta aaggcagagg 1140  
 42 ctgcttaca ctatgctcct tcatgggact cggccttcca gttgcagaat caagatccta 1200  
 43 agctaggaag tcagcagcac atggcgcaga cacccccaga aattaattcc gggcttgatt 1260  
 44 ttttgcagcc aggaagacaa gaggactcct attttacttt gggatgacta gactttcttg 1320  
 45 atttgggtgc tgaattggga ttgtaaatcc gaagttgttg aagctaaaag cggcgactat 1380  
 46 gaaactggaa ttttggaaac gcttattggt cctgggtgtt gtcttagttc tagtctgttt 1440  
 47 atgtactaga acttgacata taggaggctt ttgaaagctg aacaaacgaa gtgtgaatta 1500  
 48 ttttcttttt ttgtttttct gcagcgatgt atactaacat ctctactact aaaattacgt 1560  
 49 ctcttcgtct tactaacag tagggtggag ctgattctct tttaagtttt tcagaagggg 1620  
 50 aattcagcta tgagttttaga ggcagggtag tgtagttcag tgagcagatt ctttctgtag 1680  
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 52 tggaggttgt atct 1754  
 55 <210> SEQ ID NO: 2  
 56 <211> LENGTH: 885  
 57 <212> TYPE: DNA  
 58 <213> ORGANISM: Catharanthus roseus

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60 &lt;400&gt; SEQUENCE: 2

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61 caacaataat gtatcaatca aatgcccata attccgatca tctaacccttc ttaccacctt 60
62 tagtagatta tcaattcctc aacaacgatt ttgatttttc agaaatattt acagatttca 120
63 attacgctaa ttataattat aatacttcta cctcagataa tttctctggg tttcaattca 180
64 atgaaaattg cgaagaaaatt atttcaccaa attatgcttc ggaagattta tcggatatta 240
65 ttttaacaga tatttttcaag gatcaggata attacgaaga cgaagtcgtt gcgggagaac 300
66 aagaagaaga attaattacg acacctacct ctccgcgcgcg cggcggcggc ggatgtgagc 360
67 agagatcgaa tgaggaatgg attaggtacc gtggcgtag acggcggcca tgggggaaat 420
68 tcgctgcgga aatcagggat cccaagagaa aaggatcgag gatatgggtg ggaacttacg 480
69 agacggcgga agatgcggca ttagctttcg atcaagcggc gtttcaactc cgtggttcta 540
70 gagctagggt aaattttccc aatcttattg gttctgctaa tgctccgggt agagtaagtc 600
71 ctacagcccg atcttcatcg tgtcatcttc gtctcaata atcctatcca cagttccatg 660
72 gggatagtaa attttttctt tgagttttt agaagttata ttatctattg aaaaaataca 720
73 aaacattgca aatatTTTTT tagtacgtct ctatacttct ttttagtaat attcggatca 780
74 tgagcatggg gaagggtgata ttatccattg tcataaatta atagatacac tatcataaat 840
75 taatatgtac gaattacaag taaaatatag taagtgttaa tattg 885

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78 &lt;210&gt; SEQ ID NO: 3

79 &lt;211&gt; LENGTH: 792

80 &lt;212&gt; TYPE: DNA

81 &lt;213&gt; ORGANISM: Catharanthus roseus

83 &lt;400&gt; SEQUENCE: 3

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84 ttctaaaaaa gaagaaaaat gtccgaagaa atcatttccg tctcagatcg atttcttctt 60
85 tccttaaatcg aagaacatct tctcagcgat aattctgatg attccagctc ggaattgact 120
86 tctacagagg aaaattggga agaaattttt gcagattttc taaattgggc gggatccgaa 180
87 atacagaaac gcggtagccc gagttccgaa agctgtcaat cgaattcaat ggcggaaagc 240
88 tgtcaggagg attctgttgt gggaaccccc ccagaagcgg cggccggagg aggttggtcg 300
89 aaggattgga accggtataa gggcgtaga cggcggccgt gggggaagtt cgcggcggag 360
90 ataagggatc cgaaaaagaa aggatccagg atttggttg gtacatacga gacacctgag 420
91 gatgcagcat tggttatga tgcagccgcg tttaatatgc gtggagctaa agctaggctt 480
92 aattttcctc atttgattgg ttccaatatt tccggaccog ttagagtaaa cccgagaaaa 540
93 cgtttccctg cggagccttc tacgacgtcg tcgtcttctt cttcttcttc gtctgaaaaa 600
94 agtggaggaa ggaagaagag acgatattaa ttaattatta aaagtggagg attaaaaaaa 660
95 ttctgtgaaa tgagagatta ttacgtgggt tttgttaagc ccgataatcc ctcatgttaa 720
96 aattattaac ttcacgatg ttctttttta aatctttgga atgtacaaaa ttttatatcc 780
97 aaaaaagttc ac 792

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100 &lt;210&gt; SEQ ID NO: 4

101 &lt;211&gt; LENGTH: 376

102 &lt;212&gt; TYPE: PRT

103 &lt;213&gt; ORGANISM: Catharanthus roseus

105 &lt;400&gt; SEQUENCE: 4

```

106 Met Ala Leu Leu Asp Gln Ala Ser Asn Leu Ser Pro Met Pro Phe Asp
107   1           5           10           15
109 Phe Thr Arg Lys Arg Lys Ser Arg Arg Arg Asp Gly Thr Lys Asn Val
110           20           25           30
112 Ala Glu Thr Leu Ala Lys Trp Lys Glu Tyr Asn Glu Lys Leu Asp Ala
113           35           40           45
115 Leu Asp Gly Gly Lys Pro Ala Arg Lys Val Pro Ala Lys Gly Ser Lys
116           50           55           60
118 Lys Gly Cys Met Lys Gly Lys Gly Gly Pro Glu Asn Ser His Cys Lys

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119 65                               70                               75                               80
121 Tyr Arg Gly Val Arg Gln Arg Thr Trp Gly Lys Trp Val Ala Glu Ile
122                               85                               90                               95
124 Arg Glu Pro Asn Arg Gly Ser Arg Leu Trp Leu Gly Thr Phe Arg Asn
125                               100                              105                              110
127 Ala Ile Glu Ala Ala Leu Ala Tyr Asp Glu Ala Ala Arg Ala Met Tyr
128                               115                              120                              125
130 Gly Pro Cys Ala Arg Leu Asn Leu Pro Asn Tyr Arg Ala Ser Glu Glu
131                               130                              135                              140
133 Ser Ser Ser Leu Pro Thr Thr Ser Gly Ser Asp Thr Thr Thr Ala Ser
134 145                               150                              155                              160
136 Gly Ile Ser Glu Val Ser Val Tyr Glu Asp Lys Lys Phe Thr Pro Val
137                               165                              170                              175
139 Val Ser Gly Leu Lys Gln Asp Asp Lys Gly Glu Ser Leu Glu Ser Ala
140                               180                              185                              190
142 Asp Ser Lys Pro Gln Leu Leu Val Asp Ala Gly Thr Pro Met Ser Ala
143                               195                              200                              205
145 Val Lys Glu Glu Pro Lys Glu Tyr Gln Val Met Asp Ser Gln Ser Glu
146                               210                              215                              220
148 Gly Gln Phe Gly Asp Glu Glu Pro Pro Ser Lys Leu Val Cys Lys Glu
149 225                               230                              235                              240
151 Val Asp Phe Gly Gln Asp Gln Ala Val Val Pro Ala Val Lys Asn Ala
152                               245                              250                              255
154 Glu Glu Met Gly Gly Glu Met Gly Gly Asp Ile Leu Lys Gly Cys Ser
155                               260                              265                              270
157 Leu Ser Glu Met Phe Asp Val Asp Glu Leu Leu Ser Val Leu Asp Ser
158                               275                              280                              285
160 Thr Pro Leu His Ala Ser Asp Phe Gln His Gly Met Gly Asn Gly Asn
161                               290                              295                              300
163 Val Lys Ala Glu Ala Ala Tyr Asn Tyr Ala Pro Ser Trp Asp Ser Ala
164 305                               310                              315                              320
166 Phe Gln Leu Gln Asn Gln Asp Pro Lys Leu Gly Ser Gln Gln His Met
167                               325                              330                              335
169 Ala Gln Thr Pro Pro Glu Ile Asn Ser Gly Leu Asp Phe Leu Gln Pro
170                               340                              345                              350
172 Gly Arg Gln Glu Asp Ser Tyr Phe Thr Leu Gly Asp Leu Asp Phe Leu
173                               355                              360                              365
175 Asp Leu Gly Ala Glu Leu Gly Leu
176                               370                              375
180 <210> SEQ ID NO: 5
181 <211> LENGTH: 210
182 <212> TYPE: PRT
183 <213> ORGANISM: Catharanthus roseus
185 <400> SEQUENCE: 5
186 Met Tyr Gln Ser Asn Ala His Asn Ser Asp His Leu Thr Phe Leu Pro
187 1                               5                               10                               15
189 Pro Leu Val Asp Tyr Gln Phe Leu Asn Asn Asp Phe Asp Phe Ser Glu
190                               20                               25                               30
192 Ile Phe Thr Asp Phe Asn Tyr Ala Asn Tyr Asn Tyr Asn Thr Ser Thr

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193          35          40          45
195 Ser Asp Asn Phe Ser Gly Phe Gln Phe Asn Glu Asn Cys Glu Glu Ile
196          50          55          60
198 Ile Ser Pro Asn Tyr Ala Ser Glu Asp Leu Ser Asp Ile Ile Leu Thr
199 65          70          75          80
201 Asp Ile Phe Lys Asp Gln Asp Asn Tyr Glu Asp Glu Val Val Ala Gly
202          85          90          95
204 Glu Gln Glu Glu Glu Leu Ile Thr Thr Pro Thr Ser Arg Gly Gly Gly
205          100          105          110
207 Gly Gly Gly Cys Glu Gln Arg Ser Asn Glu Glu Trp Ile Arg Tyr Arg
208          115          120          125
210 Gly Val Arg Arg Arg Pro Trp Gly Lys Phe Ala Ala Glu Ile Arg Asp
211          130          135          140
213 Pro Lys Arg Lys Gly Ser Arg Ile Trp Leu Gly Thr Tyr Glu Thr Ala
214 145          150          155          160
216 Glu Asp Ala Ala Leu Ala Phe Asp Gln Ala Ala Phe Gln Leu Arg Gly
217          165          170          175
219 Ser Arg Ala Arg Leu Asn Phe Pro Asn Leu Ile Gly Ser Ala Asn Ala
220          180          185          190
222 Pro Val Arg Val Ser Pro Arg Arg Arg Ser Ser Ser Cys His Leu Arg
223          195          200          205
225 Pro Gln
226          210
230 <210> SEQ ID NO: 6
231 <211> LENGTH: 203
232 <212> TYPE: PRT
233 <213> ORGANISM: Catharanthus roseus
235 <400> SEQUENCE: 6
236 Met Ser Glu Glu Ile Ile Ser Val Ser Asp Arg Phe Leu Leu Ser Leu
237 1          5          10          15
239 Ile Glu Glu His Leu Leu Ser Asp Asn Ser Asp Asp Ser Ser Ser Glu
240          20          25          30
242 Leu Thr Ser Thr Glu Glu Asn Trp Glu Glu Ile Phe Ala Asp Phe Leu
243          35          40          45
245 Asn Trp Ser Gly Ser Glu Ile Gln Lys Arg Gly Ser Pro Ser Ser Glu
246          50          55          60
248 Ser Cys Gln Ser Asn Ser Met Ala Glu Ser Cys Gln Glu Asp Ser Val
249 65          70          75          80
251 Val Gly Thr Pro Pro Glu Ala Ala Ala Gly Gly Gly Cys Ser Lys Asp
252          85          90          95
254 Trp Asn Arg Tyr Lys Gly Val Arg Arg Arg Pro Trp Gly Lys Phe Ala
255          100          105          110
257 Ala Glu Ile Arg Asp Pro Lys Lys Lys Gly Ser Arg Ile Trp Leu Gly
258          115          120          125
260 Thr Tyr Glu Thr Pro Glu Asp Ala Ala Leu Ala Tyr Asp Ala Ala Ala
261          130          135          140
263 Phe Asn Met Arg Gly Ala Lys Ala Arg Leu Asn Phe Pro His Leu Ile
264 145          150          155          160
266 Gly Ser Asn Ile Ser Gly Pro Val Arg Val Asn Pro Arg Lys Arg Phe

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267          165          170          175
269 Pro Ala Glu Pro Ser Thr Thr Ser Ser Ser Ser Ser Ser Ser Ser
270          180          185          190
272 Glu Asn Ser Gly Gly Arg Lys Lys Arg Arg Tyr
273          195          200
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278 <211> LENGTH: 48
279 <212> TYPE: DNA
280 <213> ORGANISM: Catharanthus roseus
282 <400> SEQUENCE: 7
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286 <210> SEQ ID NO: 8
287 <211> LENGTH: 25
288 <212> TYPE: DNA
289 <213> ORGANISM: Artificial Sequence ✓
291 <220> FEATURE:
292 <223> OTHER INFORMATION: Description of Artificial Sequence: primer ✓
294 <400> SEQUENCE: 8
295 ccacgtggtt gtagtctctt agacc          25
298 <210> SEQ ID NO: 9
299 <211> LENGTH: 25
300 <212> TYPE: DNA
301 <213> ORGANISM: Artificial Sequence ✓
303 <220> FEATURE:
304 <223> OTHER INFORMATION: Description of Artificial Sequence: primer ✓
306 <400> SEQUENCE: 9
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311 <211> LENGTH: 26
312 <212> TYPE: DNA
313 <213> ORGANISM: Artificial Sequence ✓
315 <220> FEATURE:
316 <223> OTHER INFORMATION: Description of Artificial Sequence: primer ✓
318 <400> SEQUENCE: 10
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322 <210> SEQ ID NO: 11
323 <211> LENGTH: 21
324 <212> TYPE: DNA
325 <213> ORGANISM: Artificial Sequence ✓
327 <220> FEATURE:
328 <223> OTHER INFORMATION: Description of Artificial Sequence: primer ✓
330 <400> SEQUENCE: 11
331 agaccgcgaa gaatgaaagt g          21
334 <210> SEQ ID NO: 12
335 <211> LENGTH: 29
336 <212> TYPE: DNA
337 <213> ORGANISM: Artificial Sequence ✓
339 <220> FEATURE:
340 <223> OTHER INFORMATION: Description of Artificial Sequence: primer ✓

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VERIFICATION SUMMARY

DATE: 08/14/2001

PATENT APPLICATION: US/09/890,782

TIME: 10:55:51

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Output Set: N:\CRF3\08142001\I890782.raw

L:10 M:270 C: Current Application Number differs, Replaced Current Application Number

L:11 M:271 C: Current Filing Date differs, Replaced Current Filing Date